

New Hampshire Coronavirus Disease 2019 Weekly Partner Call

October 28, 2021

Ben Chan Elizabeth Talbot Beth Daly Lindsay Pierce

Thursday noon-time partner calls will focus on science, medical, and vaccine updates with time for Q&A



Healthcare Provider & Public Health Partner Calls are Being Re-Scheduled (to Avoid Holidays)

- Moving to the 1st and 3rd Thursday of each month from 12:00-1:00 pm (Next call will be November 4th)
- Webinar/call information (stays the same):
 - Zoom link: https://nh-dhhs.zoom.us/s/94059287404
 - Webinar ID: 940 5928 7404
 - Passcode: 353809
 - Telephone: 646-558-8656



Agenda

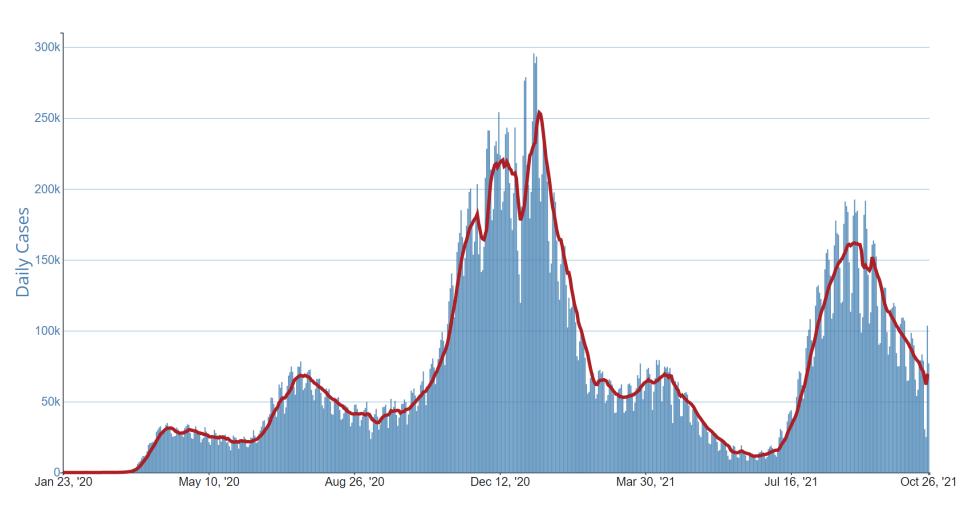
- Epidemiology Update
- CDC's Updated Recommendations for Use of COVID-19 Vaccines (Focus on Booster Doses)
- Heterologous Booster Dosing
- Questions & Answers (Q&A)



Epidemiology Update

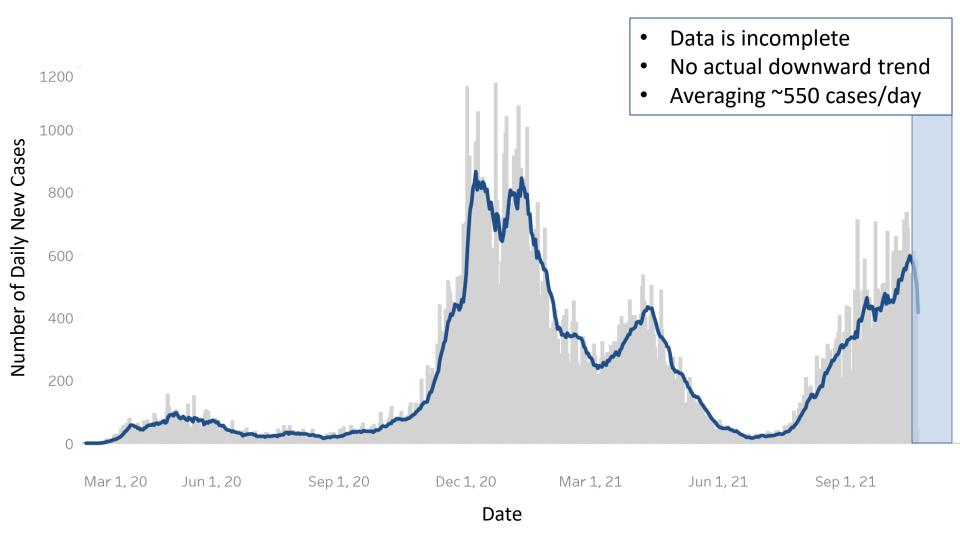


U.S. National Daily Incidence of COVID-19



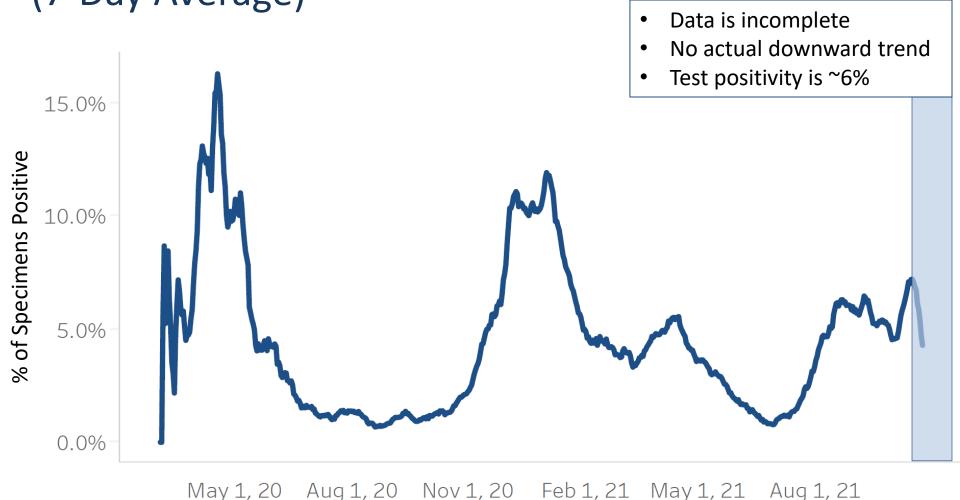


Number of New COVID-19 Cases per Day in NH





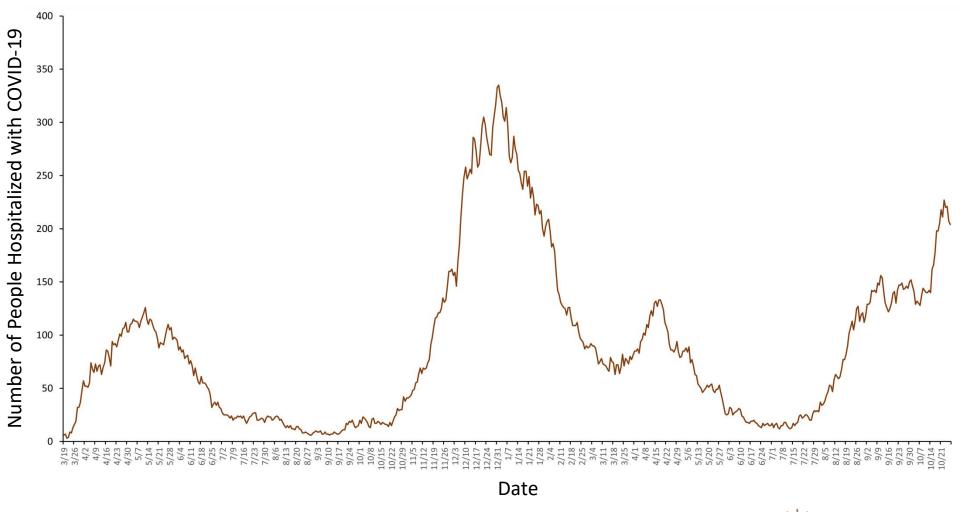
% of Tests (Antigen and PCR) Positive for COVID-19 (7-Day Average)



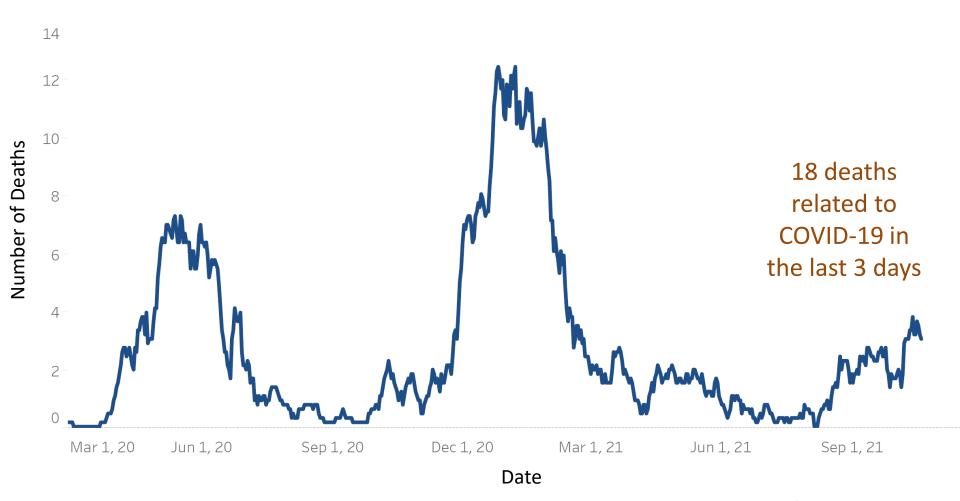
Date Laboratory Test Completed



Number of People Hospitalized with COVID-19 Each Day in NH (Hospital Census)



Average Number of COVID-19 Deaths per Day in NH (Based on Date of Death)





CDC's Updated Recommendations for Use of COVID-19 Vaccines (Booster Doses)



Review the Following

- NH DPHS <u>COVID-19 HAN, Update #47</u> about COVID-19 vaccine booster dose recommendations
- CDC's <u>Interim Clinical Considerations for Use of COVID-19</u>
 <u>Vaccines</u> (updated October 25th)



Booster Dose Update: Key Points

- Any person 18+ years of age is now eligible for a booster dose as long as the person is either:
 - At least 6 months beyond completion of a primary COVID-19 vaccine series with the Pfizer-BioNTech or Moderna vaccine (including any 3rd dose administered because a person is moderate-severely immunocompromised)

OR

- At least 2 months beyond completion of the single-dose J&J/Janssen COVID-19 vaccine
- Heterologous (mix-and-match) dosing is now allowed any of the COVID-19 vaccines can be used for a booster dose regardless of the vaccine product used for the primary series
- Recommendations on who should receive a booster dose and the timing of booster administration depends on the primary series



Recommendations Based on Primary Series

- If a person completed their primary COVID-19 vaccine series with the Janssen vaccine, a booster dose is recommended starting at least 2 months after receiving the single-dose Janssen vaccine (regardless of risk factors or underlying medical conditions)
- If a person completed their primary COVID-19 vaccine series with the Pfizer-BioNTech or Moderna vaccine, a booster dose is permitted or recommended (based on age and risk factors – see next slide) starting at least 6 months after completing a primary mRNA COVID-19 vaccine series



Recommendations for Persons Who Completed a Primary Series with the Pfizer-BioNTech or Moderna COVID-19 Vaccine

Persons who **should** receive a booster dose:

- 65 years of age or older
- 18 years of age or older who reside in a long-term care setting
- 50-64 years of age with certain underlying medical conditions

Persons who **may** receive a booster dose:

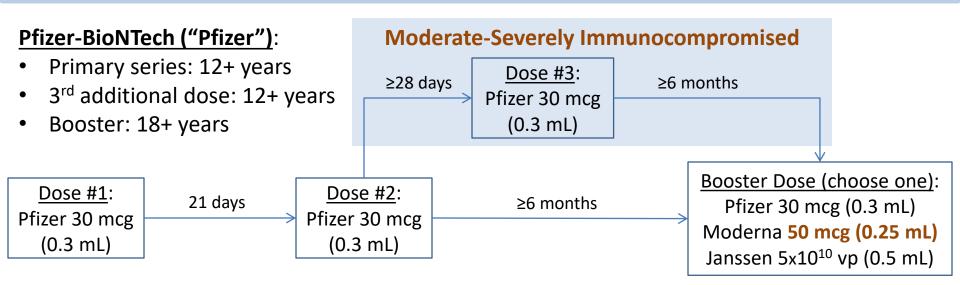
- 18-49 years of age with certain underlying medical conditions
- 18-64 years of age who are at <u>increased</u> <u>risk</u> for SARS-CoV-2 exposure and transmission because of an occupational or institutional setting
- The "may receive" recommendation is based on an individual's own assessment of their risks/benefits from a booster
- Any of the following COVID-19 vaccines can be used as a booster: Pfizer-BioNTech, Moderna, or J&J/Janssen
- The Moderna booster dose is a half-dose (50 mcg)

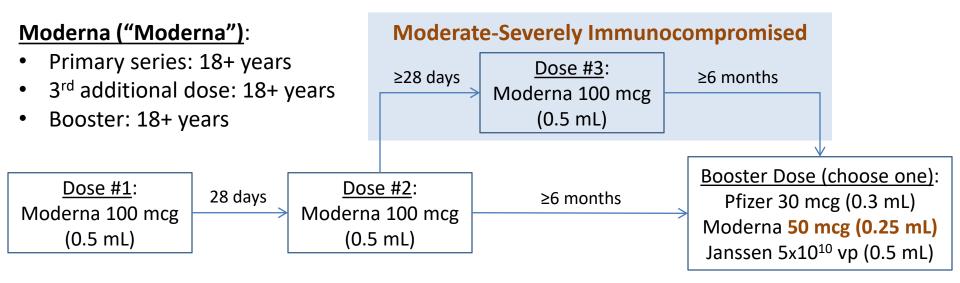


CDC's Summary Table of Vaccine Recommendations (Primary Series & Booster)

Vaccine		Primary s	Booster dose			
	Dose (volume)	No. doses (interval)	Age (yrs)	Interval from primary to booster dose	Dose (volume)	Age (yrs)
				booster dose		
Pfizer-	30 μg	2	≥12	≥6 months	30 μg	≥18
BioNTech	(0.3 ml)	(21 days)			(0.3 ml)	
Moderna	100 μg	2	≥18	≥6 months	50 μg	≥18
	(0.5 ml)	(28 days)			(0.25 ml)	
Janssen	$5 \times 10^{10} \text{ VP}$	1	≥18	≥2 months	$5 \times 10^{10} \text{ VP}$	≥18
	(0.5 ml)	(N/A)			(0.5 ml)	

- Table does not include recommendations for a 3rd additional mRNA vaccine dose for people who are moderate-severely immunocompromised
- Any of the COVID-19 vaccines (Pfizer-BioNTech, Moderna, Janssen) can be used for booster vaccination, regardless of the vaccine product used for primary vaccination







Johnson & Johnson/Janssen ("J&J/Janssen"):

- Primary "series" (single dose): 18+ years
- Additional dose for moderate-severely immunocompromised: N/A
- Booster: 18+ years



For persons who completed their primary vaccine series with the single-dose Janssen vaccine, studies are showing higher antibody boosting when the Pfizer or Moderna vaccines are used for a booster...



Heterologous Booster Dosing



Mixing Vaccine Presentations

Are heterologous prime-boost schedules safe and effective?

Heterologous vs Homologous Boosters

- Would ease logistical problems inherent in vaccine programs
- May induce enhanced or more durable immune response

CDC Oct 21 2021

- The same product that was used for the primary regimen should be used for the booster
 - If that is not available or another product is preferred, heterologous boosting with a single dose of any of the authorized COVID-19 vaccine boosters is acceptable
- Heterologous dosing may be considered for the booster dose only
 - All doses of the primary series and additional dose (if indicated for moderately to severely immunocompromised people who received 2 doses of mRNA vaccine) should utilize the same vaccine product
- Individual benefit-risk assessment may inform which booster product to use
 - Availability of booster product
 - Risk profile of vaccine boosters, including rare events



Additional Detailing Re Interval

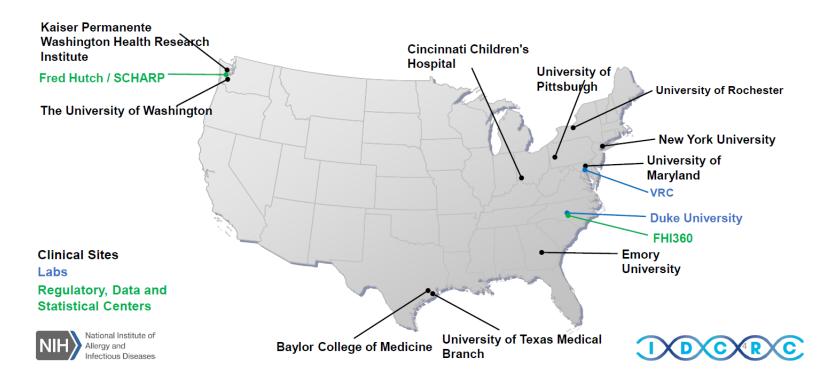
- Interval should follow the interval recommended by the primary series
 - People who received a single dose Janssen primary series can receive a mRNA COVID-19 booster dose at least 2 months after completing primary series
 - If Moderna vaccine booster is used, the booster dose and volume should be utilized (50μg in 0.25ml)

Vaccine	Primary series/dose			Additional dose in immunocompromised *			Booster dose+		
	Dose (volume)	N doses (interval)	Age (y)	Dose (volume)	N doses (interval since primary series)	Age (y)	Dose (volume)	N doses (interval since primary series#	Age (y)
Pfizer- BioNTech	30 μg (0.3 ml)	2 (21 d)	≥12	30 μg (0.3 ml)	1 (≥28 d)	≥12	30 μg (0.3 ml)	1 (≥6 m)	≥18§
Moderna	100 μg (0.5 ml)	2 (28 d)	≥18	100 μg (0.5 ml)	1 (≥28 d)	≥18	50 μg (0.25 ml)	1 (≥6 m)	≥18§
Janssen	5 × 10 ¹⁰ VP (0.5 ml)	1 (N/A)	≥18	(see booster dose)			5 × 10 ¹⁰ VP (0.5 ml)	1 (≥2 m)	≥18



VRBBPAC Oct 14-15

The "Mix and Match" Study Team Co-Chairs: Kirsten E. Lyke, MD and Robert L. Atmar, MD





Design

- Phase 1/2 volunteers >18y received EUA approved vaccine as booster (Moderna at 100ug dose) at least 12w after primary
- Studied safety, immunogenicity and reactogenicity

	Group	Sample Size*	EUA Vaccine	Interval Delayed Booster (weeks) Vaccination		Strategy Tested		
Γ	1	50	Previously dosed Janssen – Ad26.COV2-S	≥12	Moderna- mRNA-1273	Same Strain Heterologous platform		
Moderna (100 mcg)	2	50	Previously dosed Moderna – mRNA-1273	≥12	Moderna- mRNA-1273	Control - Same Strain & platform		
	3	50	Previously dosed Pfizer/BioNTech -BNT162b2	≥12	Moderna- mRNA-1273	Same Strain Similar platform		
(100 mcg)	4	50	Previously dosed Janssen – Ad26.COV2-S	≥12	Janssen – Ad26.COV2.S	Control - Same Strain & platform		
Janssen (5x1010 vp)	5	50	Previously dosed Moderna – mRNA-1273	≥12	Janssen – Ad26.COV2.S	Same Strain Heterologous platform		
	6	50	Previously dosed Pfizer/BioNTech -BNT162b2	≥12	Janssen – Ad26.COV2.S	Same Strain Heterologous platform		
Boostel	7	50	Previously dosed Janssen – Ad26.COV2-S	≥12	Pfizer/BioNTech – BNT162b2	Same Strain Heterologous platform		
Pfizer (30 mcg)	8	50	Previously dosed Moderna – mRNA-1273	≥12	Pfizer/BioNTech- BNT162b2	Same Strain Similar platform		
(30 meg)	9	50	Previously dosed Pfizer/BioNTech -BNT162b2	≥12	Pfizer/BioNTech – BNT162b2	Control - Same Strain & platform		

Study Visits: Days 1, 8 (call), 15, 29, Months 3, 6, 12

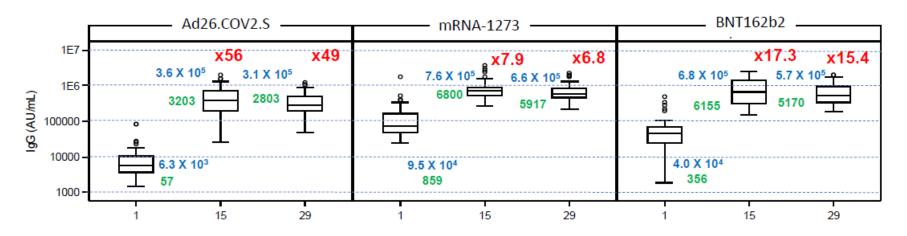


N=458

Table 1. Characteristics of the Participants at Enrollment										
Group	1	2	3	4	5	6	7	8	9	
Primary	Janssen	Moderna	Pfizer/BioNTech	Janssen	Moderna	Pfizer/BioNTech	Janssen	Moderna	Pfizer/BioNTech BNT162b2	
EUA Immunization Vaccine	Ad26.COV2-S	mRNA-1273	BNT162b2	Ad26.COV2-S	mRNA-1273	BNT162b2	Ad26.COV2-S	mRNA-1273	30-mcg	
	5x10 ¹⁰ vp	100-mcg	30-mcg	5x1010 vp	100-mcg	30-mcg	5x1010 vp	100-mcg		
Booster	Moderna mRNA-1273 100-mcg			Janssen Ad26.COV2-S 5x10 ¹⁰ vp			Pfizer/BioNTech BNT162b2 30-mcg			
Total Number	53	51	50	50	49	51	53	51	50	
Sex – no. (%)										
Female	26 (49.1)	32 (62.7)	29 (58.0)	27 (46.0)	16 (32.7)	23 (45.1)	29 (54.7)	26 (51.0)	23 (46.0)	
Male	27 (50.9)	19 (37.3)	21 (42.0)	23 (54.0)	33 (67.3)	28 (54.9)	24 (45.3)	25 (49.0)	27 (54.0)	
Age – years										
Mean (s.d.)	56.8 (14.5)	53.1 (16.2)	54.8 (17.4)	50.1 (13.9)	49.9 (16.8)	50.3 (15.4)	47.7 (14.5)	54.3 (16.8)	50.4 (17.9)	
Range	24-81	24-76	22-85	24-77	20-75	20-76	22-74	23-75	19-80	
Race – no. (%)										
Asian	4 (7.5)	5 (9.8)	4 (8.0)	3 (6.0)	5 (10.2)	6 (11.8)	1 (1.9)	2 (3.9)	1 (2.0)	
Hawalian or Pacific Islander	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (1.9)	0 (0.0)	0 (0.0)	
Black/African American	1 (1.9)	2 (3.9)	3 (6.0)	0 (0.0)	0 (0.0)	2(3.9)	0 (0.0)	2 (3.9)	1 (2.0)	
White	46 (86.8)	41 (80.4)	43 (86.0)	44 (88.0)	43 (87.8)	40 (78.4)	50(94.3)	47 (92.2)	43 (86.0)	
Multi-racial	1 (1.9)	3 (5.9)	0 (0.0)	3 (6.0)	1 (2.0)	2 (3.9)	1 (1.9)	0 (0.0)	4 (8.0)	
Other	1 (1.9%)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (2.0%)	0 (0.0)	0 (0.0)	1 (2.0%)	
Ethnicity – no (%)										
Non-Hispanic	49 (92.5)	46 (90.2)	47 (94.0)	47 (94.0)	49 (100.0)	48 (94.1)	51 (96.2)	49 (96.1)	45 (90.0)	
Hispanic/Latino	4 (7.5)	4 (7.8)	3 (6.0)	2 (4.0)	0 (0.0)	3 (5.9)	2 (3.8)	2 (3.9)	5 (10.0)	
Unknown/Not reported	0 (0.0)	1 (2.0)	0 (0.0)	1 (2.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	
Boost Interval weeks										
Mean (s.d.)	13.7 (1.0)	16.4 (1.9)	16.8 (2.2)	17.7 (2.0)	19.3 (4.2)	20.6 (5.8)	19.9 (2.5)	22.9 (4.6)	24.1 (5.2)	
Range	12.0-15.9	12.4-20.0	12.0-20.9	13.9-21.0	12.6-26.0	12.3-41.3	10.9-23.0	12.6-28.7	14.3-31.9	



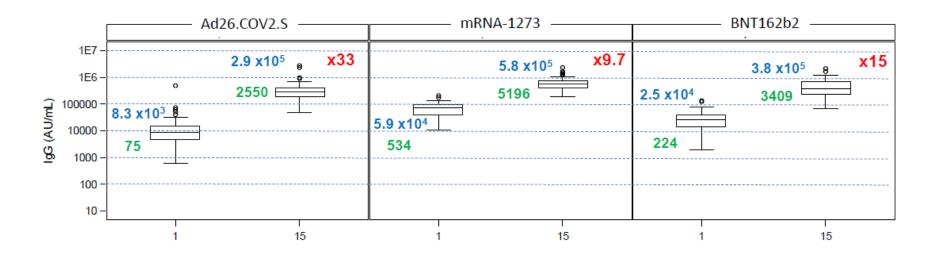
Moderna As Booster: IgG



Appeared similar in age groups 18-55 and 56+ NA Ab to Spike D614G showed similar patterns



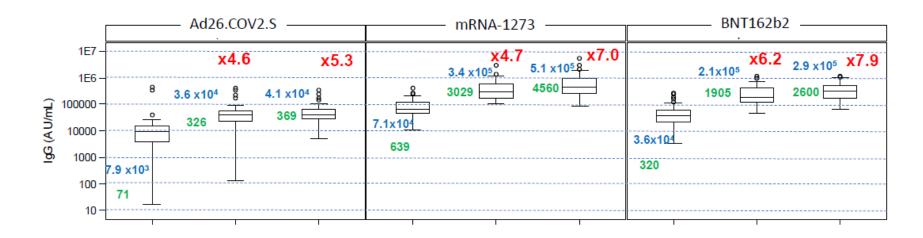
Pfizer As Booster: IgG



Appeared similar in age groups 18-55 and 56+ NA Ab to Spike D614G showed similar patterns

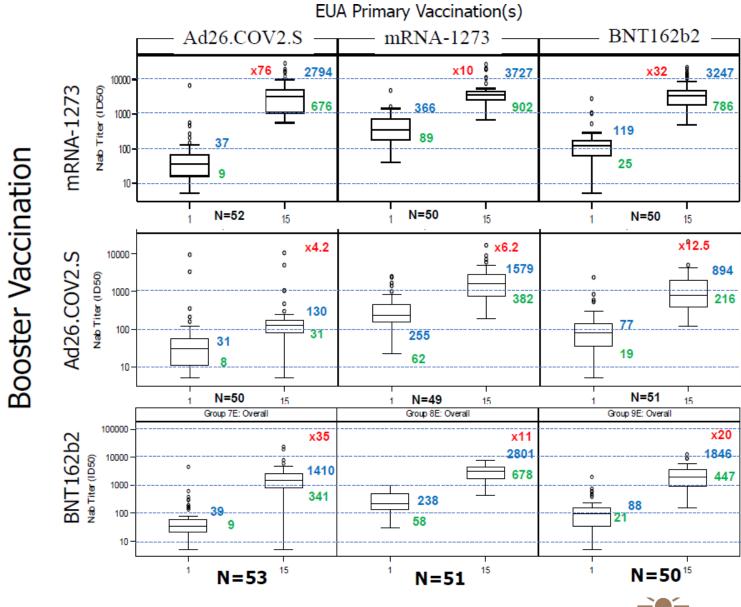


Janssen As Booster: IgG



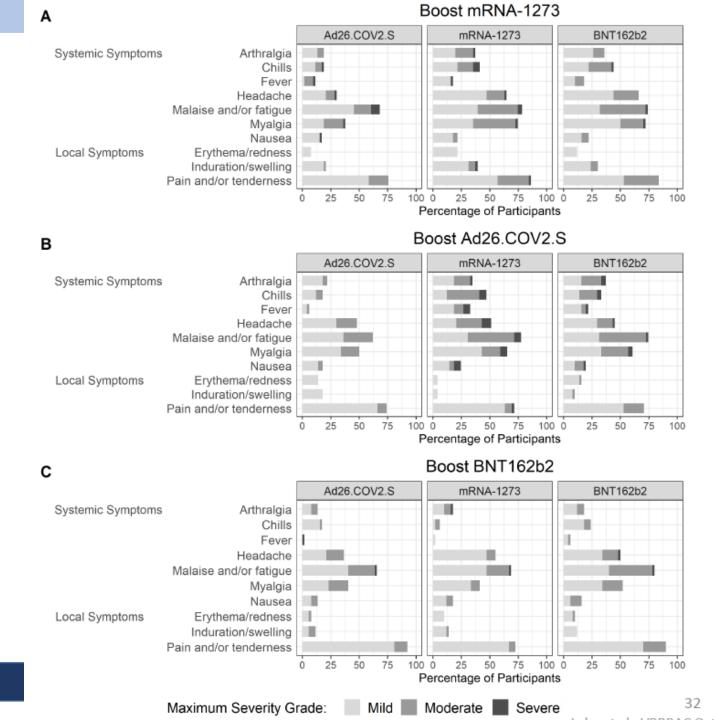
Appeared similar in age groups 18-55 and 56+







Solicited Adverse Reactions At D8



Safety

Unsolicited AEs (deemed related to boost) of any severity grade

- mRNA-1273: 24/154 (15.6%)
- Ad26.COV2.S: 18/150 (12.0%)
- BNT162b2: 22/154 (14.3%)

Most related AEs were Grade 1 or 2 severity

Four related Grade 3 AEs:

- Vomiting in one participant mRNA-1273 booster group
- Vomiting in one participant Ad26.COV2.S booster group
- Fatigue in one participant Ad26.COV2.S booster group
- Insomnia in one participant Ad26.COV2.S booster group



Limitations

- Non-randomized, open label design
- Study not designed to compare between boosts
 - Didn't control for intervals between primary vaccine and boosts
- Correlates of protection are not completely elucidated.
- Correlates for severe disease and death are even less well understood.
- This is only antibody data.
 - Cellular immune responses are still being analyzed
- These data represent only early timepoints from the trial
 - Vaccines may differ in time to reach peak responses, and may have different durability of the responses
- Healthy volunteers, short follow up
- Used 100ug Moderna boost dose



Conclusions

- 1. Use of mRNA-1273, Ad26.COV2.S and BNT162b2 as booster vaccines led to anamnestic serologic responses in all 3 EUA-dose vaccine groups
- 2. For a given primary EUA Covid-19 vaccine, heterologous boosts elicited similar or higher serologic responses as compared to their respective homologous booster responses
- 3. mRNA vaccines resulted in higher antibody titers in the first 28 days after the boost
- 4. No safety concerns identified

If you got Janssen, 15 days after boost with:

- Moderna increased Ab levels 76-fold in 15d following
- Pfizer increased Ab levels 35-fold in 15d following
- Janssen raised antibody levels 4-fold in 15d following



Singapore Study of Heterologous Boost







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Search

Comment on this pape

Differential immunogenicity of homologous versus heterologous boost in Ad26.COV2.S vaccine recipients

Nicholas Khoo Kim Huat, Joey Ming Er Lim, Upkar S. Gill, Ruklanthi de Alwis, Nicole Tan, Justin Zhen Nan Toh, Jane E. Abbott, Carla Usai,
Eng Eong Ooi, Jenny Guek Hong Low,
Nina Le Bert, Patrick T. F. Kennedy,
Antonio Bertoletti

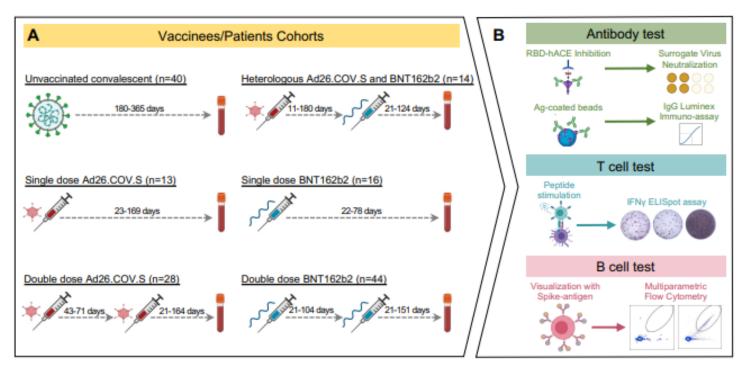
doi: https://doi.org/10.1101/2021.10.14.21264981

This article is a preprint and has not been peer-reviewed [what does this mean?]. It reports new medical research that has yet to be evaluated and so should not be used to guide clinical practice.

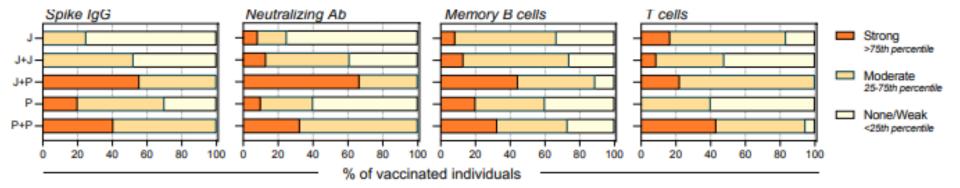


Small Singapore Study

 Strategy of boosting Ad26.COV2.S with heterologous was superior to homologous









Cov-Boost Heterologous Booster Study

- UK study at 18 sites to inform UK's booster program
 - 7 vaccines as booster doses for people >30yo who are already fully vaccinated with 2 doses of different authorized vaccine
 - AztraZeneca, Pfizer,
 Moderna, Novavax, Valneva,
 Curevac, Janssen plus control
 formulation
- May 19 began enrolling
 - Initial safety and immunologic results September
 - Follow up 1y

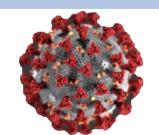
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